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REVIEW

Respiratory muscle ultrasonography: methodology, basic and advanced principles and clinical applications in ICU and ED patients—a narrative review

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Abstract

Respiratory muscle ultrasound is used to evaluate the anatomy and function of the respiratory muscle pump. It is a safe, repeatable, accurate, and non-invasive bedside technique that can be successfully applied in different settings, including general intensive care and the emergency department. Mastery of this technique allows the intensivist to rapidly diagnose and assess respiratory muscle dysfunction in critically ill patients and in patients with unexplained dyspnea. Furthermore, it can be used to assess patient–ventilator interaction and weaning failure in critically ill patients. This paper provides an overview of the basic and advanced principles underlying respiratory muscle ultrasound with an emphasis on the diaphragm. We review different ultrasound techniques useful for monitoring of the respiratory muscle pump and possible therapeutic consequences. Ideally, respiratory muscle ultrasound is used in conjunction with other components of critical care ultrasound to obtain a comprehensive evaluation of the critically ill patient. We propose the ABCDE-ultrasound approach, a systematic ultrasound evaluation of the heart, lungs and respiratory muscle pump, in patients with weaning failure.

Keywords: Ultrasonography, Diaphragm ultrasound, Respiratory muscle ultrasound, Diaphragm dysfunction

Introduction

Ultrasound imaging has become increasingly popular for the diagnosis of deranged physiology and to guide treatment in critically ill patients. In particular, the sonographic evaluation of the heart and lungs is well established and widely implemented in the intensive care unit (ICU) [1, 2]. The use of ultrasound to evaluate the respiratory muscle pump function is relatively new. The relative infrequency with which respiratory muscle ultrasound is employed in clinical practice may be attributable to the complexity of this pump (especially the number of muscles involved), the difficulty in obtaining adequate ultrasound windows, and the common assumption that ultrasound evaluation of the respiratory muscles would not alter patient management in the ICU. The aim of this review is to provide an overview of the principles and current applications of respiratory muscle ultrasound and to discuss its limitations and to describe innovative ultrasound-based techniques. We propose a systematic ultrasound-based evaluation of the respiratory muscle pump, integrated with cardiac and lung ultrasound, in ICU patients.

Anatomy of the respiratory muscles

The main muscle of inspiration is the diaphragm, a thin dome-shaped muscle positioned between the chest and

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abdomen. Contraction of the zone of apposition (cylindroid part of the diaphragm attached to the thoracic outlet) results in caudal movement of the diaphragm dome, and increases intrathoracic volume. When the load imposed on the diaphragm increases, the accessory inspiratory muscles (parasternal-, external intercostal-, scalene-, and sternocleidomastoid muscles) are recruited. With further loading, the expiratory muscles are activated to assist expiration [3]. The most prominent expiratory muscles include the transversus abdominis muscle and the internal- and external oblique muscles. The differential diagnosis of diaphragm weakness is extensive (Table E1).

Techniques and views

Diaphragm

Two ultrasound approaches to visualize the diaphragm should be performed: the mid-axillary intercostal approach at the zone of apposition, and the subcostal approach using the liver or spleen as an acoustic window. Tips and tricks of respiratory muscle ultrasound are summarized in E-Figure 1.

Intercostal approach: thickness and thickening fraction

The intercostal approach is performed with a 10–15-MHz linear array transducer positioned in a cranio-caudal direction and perpendicular to the skin in the zone of apposition between the mid-axillary or antero-axillary line, in the 8th to 11th intercostal space (Fig. 1) [4, 5]. The diaphragm appears at a depth of two to four centimeters as a three-layered structure between the pleural and peritoneal membrane (Fig. 1). Characteristically, a white linear structure is seen in the middle of the diaphragm [4, 5]. We recommend measuring diaphragm thickness perpendicular to its fiber direction between the pleural and peritoneal membrane, but not including the membranes (Fig. 1). The lower limit for normal diaphragm thickness is around 1.5 mm in healthy subjects. Reference values are given in Table 1. Diaphragm thickness is affected by body composition and gender [6, 7].

The diaphragm thickens with active shortening and, therefore, thickening fraction (TF) reflects contractile activity [8, 9]. Thickening fraction of the diaphragm (TFdi) is calculated in B-mode or M-mode as the percentage inspiratory increase in diaphragm thickness relative to end-expiratory thickness during tidal breathing (TFdi) or maximal inspiratory effort (TFdi(max)): TFdi = (end-inspiratory thickness − end-expiratory thickness)/end-expiratory thickness × 100% (Fig. 1). Inspiratory thickening of the diaphragm can be used to assess muscle function. Reference values are given in Table 1 and demonstrate a relative wide range in healthy subjects [6, 9–11]. A reasonable relationship exists between TFdi and the pressure (or electrical activity) developed by the diaphragm during unassisted breathing [9, 12] and mechanical ventilation [13]. Few studies directly evaluated the correlation between TFdi and muscle pressure (Pmus) [14, 15] and, therefore, care should be taken when estimating Pmus from TFdi.

Subcostal approach: excursion

Diaphragmatic excursion is measured with a low frequency phased-array or curved-array ("abdominal") probe (2–5 MHz) positioned just below the costal arch at the midclavicular line, with the patient in semi-seated position and by angling the ultrasound beam as much as possible cranially and perpendicular to the diaphragmatic dome (Fig. 1). The diaphragm is identified as a bright line covering the liver and the spleen. Obtaining a clear image of the left hemidiaphragm can be difficult due to the poor acoustic window of the spleen. During inspiration, the diaphragm should move toward the probe (Fig. 1). Excursion is quantified in M-mode, with the M-line placed perpendicular to the direction of motion (Fig. 1); the sweep speed is best adjusted to around 10 mm/s to obtain a minimum of three respiratory cycles within one image. Diaphragm excursion should only be measured during unassisted breathing (i.e., T-piece or minimum tolerable CPAP level), since active contraction of the diaphragm cannot be distinguished from passive displacement due to ventilator inspiratory pressures [4, 12].

In a cooperative patient, a maximum inspiratory effort is performed to assess maximal excursion. Excursion of both hemidiaphragms is compared to identify unilateral weakness or paralysis (for reference values, see Table 1). The success rate for visualizing excursion is high during tidal breathing (> 95%), whereas during maximal breathing visualization is more difficult, especially on the left side [17].

If experiencing difficulties visualizing the diaphragm from the subcostal window, movement of the liver or spleen during tidal respiration can be used as an alternative. To this end, an intercostal window at the zone of apposition in B- or M-mode is advised, using a low frequency probe [18, 19]. As there is some inconsistency in agreement between diaphragmatic- and subdiaphragmatic excursion [20, 21], it is advised to use this approach for a qualitative rather than a quantitative assessment of diaphragm motion.

Under normal conditions, expiration largely depends on the elastic recoil pressure of the respiratory system, although some diaphragm activity in early expiration has been demonstrated [22]. Nevertheless, diaphragm relaxation rate derived from excursion should not be used as a measure for diaphragm function [16].

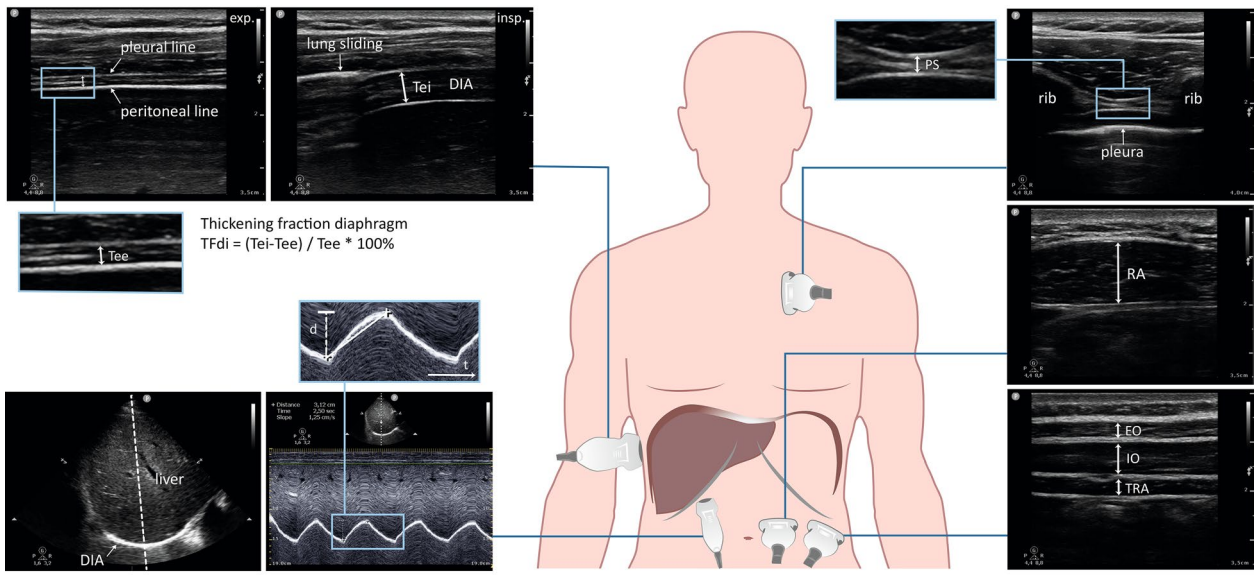


Fig. 1 Clinical application of respiratory muscle ultrasound: techniques and views

Extra-diaphragmatic inspiratory muscles

Ultrasound assessment of accessory inspiratory muscles could add information regarding patient's inspiratory effort and patient–ventilator interaction. Parasternal intercostal muscle ultrasound is performed with a 10–15 MHz linear probe positioned in cranio-caudal direction at the second intercostal space (Fig. 1). Thickness and inspiratory thickening fraction can be assessed. In healthy subjects, thickening of parasternal intercostal muscles is observed only during maximal efforts [23] and preliminary findings in ICU patients suggest the existence of a dose–response relationship between respiratory load and the parasternal intercostal thickening fraction [24].

Although the topic of future studies, parasternal intercostal muscle ultrasound may be a useful tool in the evaluation of the capacity/load balance of the respiratory muscle pump in the ventilated patient. Reference values need to be determined.

Abdominal wall expiratory muscles

Using a 10–15 MHz linear probe positioned perpendicular to the abdominal wall, with the patient in supine position, the different expiratory muscles are relatively easily to visualize as hypoechogenic layers enclosed by fascial sheaths (Fig. 1). The pressure applied to the probe should be kept to a minimum to prevent compression of the abdominal wall as this may alter the shape/thickness of the underlying muscles.

To visualize the rectus abdominis muscle, the transducer is positioned in a transverse orientation approximately 2–3 cm above the umbilicus and 2–3 cm lateral from the midline (Fig. 1) [25]. Maximum muscle thickness is obtained by sliding the probe in a cranial–caudal direction, while keeping the probe perpendicular to the skin. Next, the probe is moved laterally; the semilunar line is first identified as a thick echogenic fascia that blends lateral to the rectus abdominis muscle and medial to the oblique muscles. The external oblique, internal oblique and transversus abdominis muscles can be identified as three parallel layers (Fig. 1), usually best visualized on the anterior axillary line, midway between the inferior border of the rib cage and the iliac crest [25–27]. Reference values are provided in E-Table 2 [25–28].

Thickening fraction of the expiratory abdominal muscles (TFabd) can be calculated as the magnitude of thickness increase during expiration (TFadb = (end-expiratory thickness − end-inspiratory thickness)/end-inspiratory thickness) × 100%) and may reflect expiratory muscle effort. Preliminary data seem to demonstrate reasonable correlation between TFabd and expiratory force generation [29]. It should be noted that expiratory muscles have more degrees of freedom compared to the diaphragm; active contraction of one muscle layer may directly affect the shortening and position of the adjacent layer, which may make interpretation of TFabd more complex. Moreover, the relationship between shortening, thickening, and pressure generation is complex because of the geometry of the abdominal muscles during contraction (a

Table 1 Reference values for diaphragm muscle ultrasound in ICU patients and in the general population

Setting	Parameter	Patient position	Reference values	Abnormal values/values related to outcome	References
ICU	Thickness (mm)	–	2.4 ± 0.8		[9]
		Semi-recum	2.4 (2.0–2.9)		[75]
		Semi-recum		< 1.7	[60]
		Semi-recum	1.9 ± 0.4		[76]
	TFdi	Semi-recum		< 30%	[60]
	TFdi(max)	Semi-recum		< 36%	[59]
	TFdi	Semi-recum		< 34%	[52]
	Tidal excursion (mm)	Supine		< 11 (organ exc.)	[77]
		Semi-recum		Right < 14	[51]
	Maximal breath (mm)	Semi-recum		Left < 12	[52]
		Semi-recum		< 10	[53]
				< 25	
General population	Thickness (mm)	Sitting	1.7 ± 0.2		[78]
		Standing	2.8 ± 0.4	< 1.9	[10]
		Supine	3.3 ± 1.0	< 1.4	[6]
		Supine	1.6 ± 0.4	< 1.5	[7]
			Men: 1.9 ± 0.4	< 1.7	
			Women: 1.4 ± 0.3	< 1.3	
	TFdi(max)	Standing	37 ± 9%	< 20%	[10]
		Supine	80 ± 50%	< 20%	[6]
	Tidal excursion (mm)	Standing	Men: 18 ± 3	Men: < 10	[17]
			Women: 16 ± 3	Women: < 9	
	Sniff test (mm)	Standing	Men: 29 ± 6	Men: < 18	
			Women: 26 ± 5	Women: < 16	
	Maximal breath (mm)	Standing	Men: 70 ± 11	Men: < 47	
			Women: 57 ± 10	Women: < 37	

Thickness and excursion values are expressed in millimeter. Normal values are expressed as mean ± SD or median (range); for ICU patients, mean thickness values are reported from baseline measurements. TFdi: thickening fraction tidal breathing (or during maximal breathing = TFdi(max)), expressed in %; Position is presented as described in the original manuscript, although sitting and semi-recumbent (and potentially supine) may have overlap. Semi-recum = semi-recumbent; – = not mentioned in the manuscript

'shrinking' sphere rather than a 'shortening piston-in-a-cylinder' like the diaphragm). Future studies should confirm the relationship between expiratory muscle pressure and TFabd and its clinical relevance.

Clinical applications of respiratory muscle ultrasound (Table 2)

Role of respiratory muscle ultrasound in acute respiratory failure

Respiratory muscle weakness as the major cause for acute respiratory failure is uncommon, but should be considered if more common causes have been excluded [30, 31]. The clinical presentation of diaphragm dysfunction depends on the cause, severity and rate of progression (E-Table 1) [32]. A characteristic physical sign of bilateral diaphragm dysfunction is supine abdominal paradox: activity of accessory inspiratory muscles generates a negative inspiratory thoracic pressure (though the same pattern may be observed when respiratory loads are increased) [33]. As the diaphragm is paralyzed, this negative pressure is transferred to the abdomen, resulting in inward movement of the abdominal wall. The ultrasound corollary of this is cranial excursion of the diaphragm during inspiration, measured in M-mode. In addition, severe isolated diaphragm weakness results in an increased thickening fraction of the accessory respiratory muscles [24]. Therefore, respiratory muscle ultrasound is an excellent modality to diagnose (unilateral) diaphragm weakness or paralysis in patients with acute respiratory failure [34, 35].

Diaphragm weakness is diagnosed by an excursion of < 10–15 mm during tidal breathing or a TFdi (max) < 20% (Table 1) [6, 7, 10]. In patients with unilateral diaphragm paralysis, thickness and TFdi of the paralyzed diaphragm were significantly less compared to the other hemidiaphragm (Table 1) [10]. A left–right ratio for

thickness of < 0.5 or > 1.6 should be considered abnormal [19]. Notably, with unilateral diaphragmatic dysfunction the normal hemidiaphragm may show relatively large excursions, a compensatory mechanism to generate sufficient tidal volumes [36, 37]. For patients with bilateral paralysis, diaphragm thickness and TFdi are below reference values [10].

In patients with acute hypercapnic exacerbation of COPD (AE-COPD), diaphragm ultrasound may be used to predict success of non-invasive ventilation (NIV). Increased diaphragm excursion during NIV (> 18 mm vs < 12 mm) was associated with NIV success and a decrease in $Paco_2$ after one hour [38, 39]. Air trapping has been found to be the major limiting factor in diaphragm excursion in COPD patients [40]; therefore, improved excursion is probably an indication of reduced pulmonary hyperinflation. In a cohort of patients with AE-COPD requiring ICU admission ($n = 41$), TFdi < 20% was associated with NIV-failure ($R = 0.51$) [41], which was confirmed in a larger ($n = 75$) follow-up study (risk ratio for NIV-failure 4.4) [42]. Diaphragm ultrasound may thus reduce the risk of delayed intubation in patients with severe AE-COPD requiring NIV; however, further validation is required.

Role of respiratory muscle ultrasound in diaphragm-protective mechanical ventilation

It has been postulated that both ventilator over-assist and ventilator under-assist, resulting in muscle atrophy and muscle injury, respectively, play an important role in critical illness-associated diaphragm weakness pathophysiology [13]. To limit these detrimental consequences, it seems reasonable to titrate ventilator support such that diaphragm effort is within physiological limits, the so-called diaphragm protective mechanical ventilation [13, 43, 44]. The optimal level of diaphragm activity is currently unknown and may vary under different conditions (e.g., sepsis, weakness); however, a relatively low level of diaphragm activity, corresponding to esophageal pressure swings of 4–8 cm H_2O appears safe [45]. The role of ultrasound in diaphragm protective ventilation has not been specifically studied, but assessment of

Table 2 Clinical indications and role of ultrasound of the respiratory muscles in adults admitted in the intensive care unit or emergency department

Setting	Indication	Role of respiratory muscle ultrasound	Diagnostic performance	Limitations
ICU	Difficult weaning	Excursion and TFdi detect DD	Excursion poor to moderate TFdi moderate Better during SBT Combined with clinical parameters better performance	A significant portion of patients diagnosed with DD can be successfully extubated
	Titrate ventilator support	Detection of underuse/overuse using TFdi		Needs further validation
	Patient–ventilator interaction	Excursion and/or TFdi (compared to ventilator waveforms) can detect different types of asynchrony	Good/easy repeatable	Variability of the effectiveness between subjects Not suitable for continuous monitoring
	Estimating work of breathing	TFdi	Large range of effort at certain TFdi	Needs further validation
	Clinical suspicion of iatrogenic n. phrenicus lesion (e.g., postoperative)	Excursion can detect (unilateral) paralysis/weakness	Good	None
ED	Dyspnea of unknown origin	Excursion/TFdi can detect weakness/paralysis	High sens and spec	None
	AECOPD	Excursion/TFdi predict NIV-failure	Moderate	Needs further validation
Both	Unilateral diaphragm relaxation on chest X-ray	Excursion/TFdi/left to right ratio	Good	None
	Diagnosing and monitoring of diaphragmatic weakness/paralysis		Good, no technical failures	
	Stroke with respiratory impairment		Good detection of diaphragm involvement	None
	Neuromuscular disorders Cervical spine lesions		May help predict need for mechanical ventilation	

AECOPD acute exacerbation of chronic obstructive pulmonary disease, *DD* diaphragm dysfunction, *ED* emergency department, *ICU* intensive care unit, *NIV* non-invasive ventilation, *SBT* spontaneous breathing trial, *TFdi* diaphragm thickening fraction

TFdi, as a proxy for effort, is a reasonable approach. Data from Goligher and colleagues demonstrate that a TFdi between 15 and 30% during the first days of mechanical ventilation is associated with stable muscle thickness [44] and the shortest duration of ventilation [13]. Accordingly, a low TFdi (< 15%) in a patient on a partially supported ventilatory mode raises the possibility of ventilator overassist; therefore, decreasing assist, while monitoring other respiratory parameters (e.g., tidal volume, respiratory rate), is a reasonable approach. The upper limit for TFdi allowing diaphragm protective ventilation is more controversial. Although a moderate and statistically significant correlation exists between TFdi and diaphragm effort (Pdi, PTP) [9, 14], the range of diaphragm effort at a certain TFdi is large. We suggest that in patients with TFdi > 30–50%, ventilator support may be increased under the monitoring of other respiratory parameters to avoid hyperinflation. Given the relative imprecision of TFdi measurements, other techniques for monitoring respiratory effort should be considered.

Role of respiratory muscle ultrasound in weaning failure

An imbalance between load and capacity of the respiratory system is an important cause for SBT failure and extubation failure [46]. Therefore, respiratory muscle ultrasound could play an important role in the differential diagnosis of weaning failure. However, it is important to emphasize that a substantial proportion of patients can be successfully weaned from the ventilator despite having diaphragm dysfunction [47–49]. In addition, the clinical relevance of predicting SBT outcome with ultrasound is debatable; more relevant from a clinical perspective is the use of ultrasound to predict extubation success.

Diaphragm excursion

Kim et al. evaluated diaphragm excursion in a series of 89 patients on a T-tube, before the start of an SBT [50]. Diaphragm dysfunction–arbitrary defined as excursion < 10 mm was associated with weaning failure, but its predictive performance was poor (AUROC 0.61). Using the same cutoff, no association was found between diaphragm dysfunction and extubation failure [51]. Interestingly, when diaphragm excursion is measured after 30 min from initiation of a 2 h SBT, predictive performance of 10 mm cutoff seems to be significantly higher (AUROC 0.88) [52]. Post-cardiac surgery patients with unilateral diaphragmatic paralysis could be extubated without delay, when the contralateral diaphragm excursion was > 25 mm at maximal inspiratory effort [53]. In a meta-analysis of 10 studies evaluating diaphragm excursion to predict weaning failure and combining different definitions of weaning failure, the authors reported a sensitivity of 75% (95% CI 65–85) and specificity of 75% (95% CI 60–85), with substantial heterogeneity [54]. As diaphragm excursion is strongly dependent of lung volume [37], the reported heterogeneity could be explained by patient position and the timing of measurements, e.g., before/during the SBT and with or without ventilator assistance.

Spadaro et al. [55] evaluated the diaphragmatic-rapid shallow breathing index (D-RSBI: respiratory rate divided by diaphragm excursion) during a T-tube SBT and reported good performance as compared to RSBI alone (D-RSBI AUROC 0.89 vs RSBI AUROC 0.72, respectively, $P = 0.006$). Palkar et al. [56] evaluated the performance of the diaphragmatic excursion–time product (i.e., E–T index), the product of diaphragm excursion (cm) and inspiratory time (s). A decrease in E–T index of < 3.8% between assisted control ventilation and a PSV (5/5 H_2O) SBT had a sensitivity of 79.2% and a specificity of 75% to predict extubation success.

Remarkably, in 191 patients having successfully passed an SBT, diaphragm excursion was not associated with extubation failure [57]. This suggests that once an SBT is successfully completed, extubation outcome is primarily determined by factors other than diaphragm function.

Diaphragm thickening fraction

When performed during an SBT, TFdi > 30–36% has shown to predict extubation success [52, 54, 58]. Ferrari et al. evaluated in 46 patients ventilated through a tracheostomy tube, the role of TFdi(max) of the right hemidiaphragm during an SBT, as a predictor of weaning outcome [59] and reported that a TFdi(max) > 36% was associated with a successful SBT (sensitivity 0.82; specificity 0.88; AUROC 0.95). In another study, TFdi was calculated either during T-tube or low levels of pressure support ventilation in patients ($N = 63$) after a failed weaning attempt. A TFdi ≥ 30% had a sensitivity of 0.88 and specificity of 0.71 (AUROC 0.79) for extubation success [60]. In the above-mentioned meta-analysis assessing the predictive value of diaphragm excursions, TFdi/TFdi (max) demonstrated an AUROC of 0.87 and diagnostic odds ratio of 21 (95% CI 11–40) for prediction of weaning failure [54]. The diagnostic odds ratio is a measure of effectiveness of a diagnostic test and it is defined as the ratio of odds of a test being positive if the subject has the disease relative to the odds of the test being positive if the subject does not have the disease.

Overall, these results seem to suggest a role for diaphragm ultrasound in the differential diagnosis of patients experiencing difficult weaning, allowing bedside recognition of diaphragmatic weakness. However, the role of diaphragm ultrasound to predict successful SBT

and successful extubation requires further evaluation and currently cannot be recommended.

The ABCDE approach: a systematic ultrasound evaluation of patients with weaning failure

A weaning trial may be considered as a cardio-pulmonary stress test: it demands an increase in cardiac index, oxygen demand/consumption, and breathing effort. In most patients, post-extubation distress is the result of a combination of cardiac dysfunction, impaired gas exchange, and/or diaphragmatic dysfunction. Therefore, in patients with weaning failure we suggest the use of a structured and integrated approach that combines clinical parameters, laboratory parameters (e.g., pro-BNP) and the sonographic assessment of the lung, heart and the respiratory muscles [61]. Here, we present the ABCDE-Ultrasound approach, an intuitive aid designed to standardize the sonographic approach to weaning failure (Fig. 2). A similar but less exhaustive approach has been suggested by Mayo et al. [62]. The use of ultrasound to evaluate the heart and lung has been described in this Intensive Care Medicine critical care ultrasound series [1, 2]. Timing of ultrasound examination depends on the clinical question. For identification of patients at high-risk of weaning failure, the examination is performed before the SBT. For prediction of weaning outcome or to diagnose the cause of weaning failure, the examination is best done after the start and/or end of an SBT [62].

Role of respiratory muscle ultrasound to assess patient–ventilator interaction

Patient–ventilator asynchrony can be defined as a mismatch between the neural inspiratory time and ventilator inspiratory time [63]. Patient–ventilator asynchrony is associated with worse outcome [64] and occurs in up to half of mechanically ventilated patients. Visual inspection of the airway flow and pressure signal may detect asynchrony, but was shown to be unreliable [65]. Both measurements of esophageal pressure and diaphragm electrical activity are used as state-of-the-art techniques to assess patient–ventilator interactions [66]. These two techniques are invasive, limiting their use in daily practice. Diaphragm ultrasound might be a reasonable alternative to detect most types of patient–ventilator asynchrony (Table 3), but further studies are needed to determine its exact role [4, 67].

Limitations of respiratory muscle ultrasound

We briefly summarize current challenges of respiratory muscle ultrasound. This highlights the need for a systematic ultrasound approach to be implemented in clinical practice and research.

Reproducibility

The first and largest study regarding reliability of diaphragm thickness and TFdi measurements in mechanically ventilated patients (n = 66) was conducted by Goligher et al. [68]. Measurements were performed after marking the probe location. Intra-observer and inter-observer reproducibility coefficients for end-expiratory diaphragm thickness were 0.2 mm and 0.4 mm, respectively [68], meaning it is expected that the absolute difference between two measurements by the same observer does not differ by more than 0.2 mm on 95% of occasions (or 0.4 mm in case of two different observers). However, it should be kept in mind that 0.2 mm represents approximately 10% of total diaphragm thickness at end expiration. While in general diaphragm ultrasound seems to be a reliable technique to assess changes in diaphragm thickness over time, comparing individual patient results should be done with a degree of caution and only after adequate training, as small observer-dependent variations (e.g., measurement location, probe angulation) will affect results.

Accuracy

Accurate muscle thickness measurement depends not only on operator skills but also on technical aspects related to ultrasound physics and patient characteristics. Unclarity of surrounding membranes and insufficient ultrasound beam angulation to muscle axis may lead to measurement error. Furthermore, the spatial axial resolution (depth resolution, i.e., ½ spatial pulse length) of the probe plays a critical role. Given that the ultrasound pulse length is typically two cycles and that the ultrasound wavelength of a 10 MHz transducer is 0.15 mm (i.e., wavelength = speed of sound in soft tissue/frequency = 1540 m/s/10 MHz = 0.15 mm), the corresponding depth resolution is ½ (2 × 0.15) = 0.15 mm, which is in the same order as the intra-observer reproducibility and sufficient in order to visualize the diaphragm. Based on the ultrasound technique and equipment used, identification of the smallest detectable change that is permitted is fundamental to distinguish true changes in muscle thickness from artifact.

Research: novel techniques and future developments for functional imaging and quantification of tissue properties

Tissue Doppler imaging

Tissue Doppler imaging (TDI) quantifies the velocity of moving structures [69]. This could be an interesting modality superimposed over the B-mode diaphragm excursion images for quantification of diaphragm kinetics (video online supplement). Feasibility and reliability of diaphragm TDI have been confirmed in neonates [70] and the role of TDI for assessment of diaphragm mobility

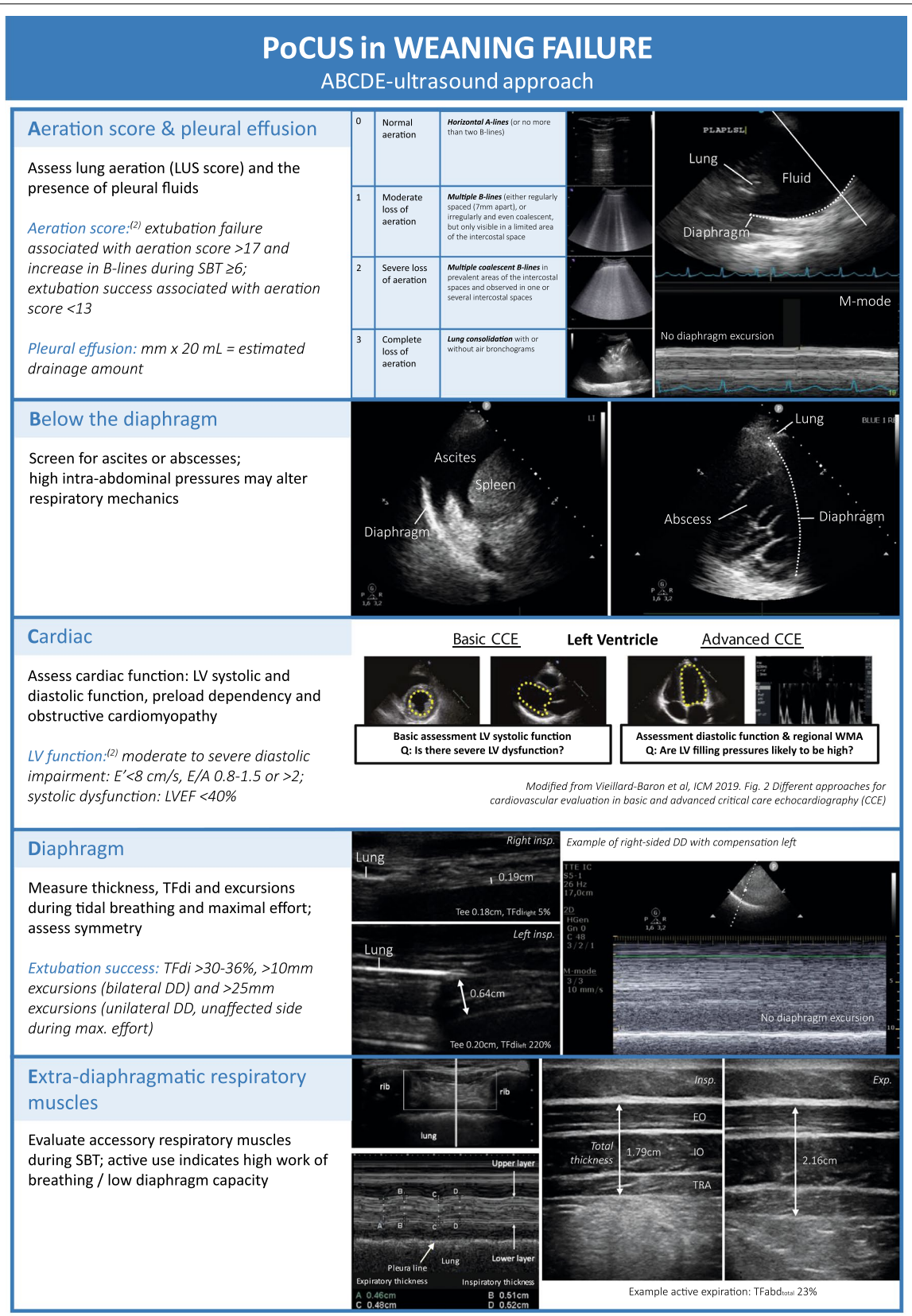
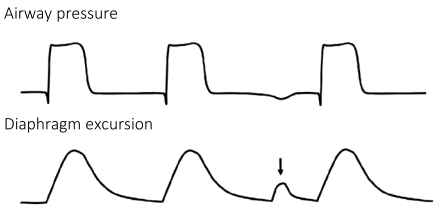
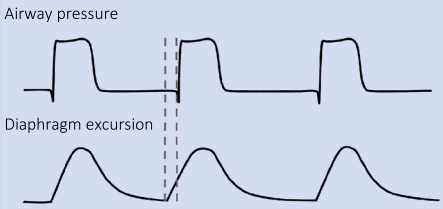
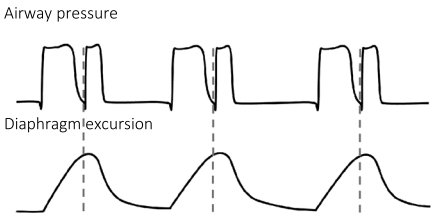
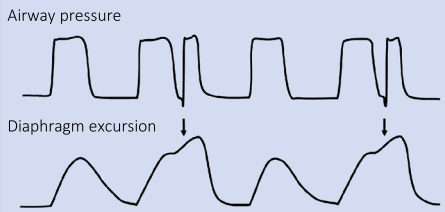
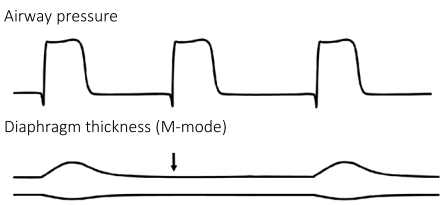

PoCUS in WEANING FAILURE
ABCDE-ultrasound approach
Aeration score & pleural effusion
Assess lung aeration (LUS score) and the presence of pleural fluids
Aeration score:(2) extubation failure associated with aeration score >17 and increase in B-lines during SBT ≥6; extubation success associated with aeration score <13
Pleural effusion: mm x 20 mL = estimated drainage amount
0 Normal aeration Horizontal A-lines (or no more than two B-lines)
1 Moderate loss of aeration Multiple B-lines (either regularly spaced (7mm apart), or irregularly and even coalescent, but only visible in a limited area of the intercostal space
2 Severe loss of aeration Multiple coalescent B-lines in prevalent areas of the intercostal spaces and observed in one or several intercostal spaces
3 Complete loss of aeration Lung consolidation with or without air bronchograms
PLAPLSL
Lung
Fluid
Diaphragm
M-mode
No diaphragm excursion
Below the diaphragm
Screen for ascites or abscesses; high intra-abdominal pressures may alter respiratory mechanics
Ascites
Spleen
Diaphragm
Lung
Abscess
Diaphragm
Cardiac
Assess cardiac function: LV systolic and diastolic function, preload dependency and obstructive cardiomyopathy
LV function:(2) moderate to severe diastolic impairment: E'<8 cm/s, E/A 0.8-1.5 or >2; systolic dysfunction: LVEF <40%
Basic CCE
Left Ventricle
Advanced CCE
Basic assessment LV systolic function
Q: Is there severe LV dysfunction?
Assessment diastolic function & regional WMA
Q: Are LV filling pressures likely to be high?
Modified from Vieillard-Baron et al, ICM 2019. Fig. 2 Different approaches for cardiovascular evaluation in basic and advanced critical care echocardiography (CCE)
Diaphragm
Measure thickness, TFdi and excursions during tidal breathing and maximal effort; assess symmetry
Extubation success: TFdi >30-36%, >10mm excursions (bilateral DD) and >25mm excursions (unilateral DD, unaffected side during max. effort)
Right insp.
Lung
0.19cm
Tee 0.18cm, TFdiright 5%
Left insp.
Lung
0.64cm
Tee 0.20cm, TFdileft 220%
Example of right-sided DD with compensation left
No diaphragm excursion
Extra-diaphragmatic respiratory muscles
Evaluate accessory respiratory muscles during SBT; active use indicates high work of breathing / low diaphragm capacity
rib
rib
lung
Upper layer
Pleura line
Lung
Lower layer
Expiratory thickness
Inspiratory thickness
A 0.46cm
B 0.51cm
C 0.48cm
D 0.52cm
Insp.
Exp.
EO
Total thickness
1.79cm
IO
TRA
2.16cm
Example active expiration: TFabdtotal 23%

Fig. 2 Point-of-Care Ultrasound (PoCUS) in patients with weaning failure: the ABCDE approach

Table 3 Types of patient–ventilator asynchronies and their ultrasound correlate

Asynchrony	Ultrasound correlate	Comments	Illustration
Wasted effort	Diaphragm excursion w/o ventilator triggering	Subcostal view M-mode Easy to perform	
Trigger delay	Delay between diaphragm excursion and ventilator triggering	Subcostal view M-mode Requires ventilator waveform display on ultrasound screen	
Double trigger	Diaphragm displacement during ventilator expiration, thereby triggering a second breath	Subcostal view M-mode During assisted modes	
Reverse trigger	Active diaphragm excursion after passive displacement, leading to a higher excursion amplitude and change in displacement slope	Subcostal view M-mode Fixed pattern Additional demonstration RT: TFdi during RT breath, no TFdi during a passive breath	
Auto-trigger	Absence of diaphragm thickening with mechanical assist	Intercostal view During assisted modes No thickening compared to other breaths	

RT reverse trigger, *TFdi* diaphragm thickening fraction

and dysfunction in patients following cardiac surgery is under investigation (ClinicalTrials.gov; NCT03295344). Potential applications include assessment of regional diaphragm contractile function at rest and with loading, and measurement of diaphragm relaxation velocity. Diaphragm relaxation abnormalities have been described as a marker of impaired contractility in patients who failed weaning [71], but nowadays they can only be assessed with invasive esophageal or diaphragmatic pressure measurements (or possibly M-mode excursion decay).

Strain imaging

Strain imaging is based on the ability to track ultrasound speckles over time and an excellent feature to quantify motion and deformation of anatomical structures. This

has a great benefit over TDI, as strain imaging is not affected by the angle between the ultrasound beam and the direction of tissue motion. Furthermore, it allows calculation of displacement, velocity, and deformation of tissue in two directions. It was recently demonstrated that strain and strain rate were highly correlated to transdiaphragmatic pressure in healthy subjects [72]. In addition, Goutman and colleagues applied speckle tracking for evaluating true diaphragm excursion in two directions, which is more accurate compared to measurements of motion along one M-mode line [73].

Shear wave elastography

Shear wave elastography is a technique that allows quantification of the elastic modulus of tissues (E-Figure. 2). Application of this technique on the diaphragm could be of clinical importance since changes in muscle stiffness may reflect alterations in muscle physiology (e.g., injury, fibrosis). Moreover, shear wave elastography is considered to be more accurate and reproducible than evaluation of echogenicity, which is highly dependent on ultrasound settings (e.g., gain, contrast, etc.). A recently published proof-of-concept work demonstrated that changes in diaphragm stiffness during inspiratory efforts as assessed with ultrasound shear wave elastography reflect changes in transdiaphragmatic pressure [74]. Therefore, it might offer a new non-invasive method for gauging diaphragm effort.

Conclusion

Respiratory muscle ultrasound is a widely available, highly feasible, non-invasive bedside radiation-free technique that can be easily applied at the bedside. It is, therefore, the imaging modality of choice for assessment of the respiratory muscles in ICU patients. Mastery of respiratory muscle ultrasound allows the intensivist to rapidly obtain information on global function of the respiratory muscle pump, in particular to diagnose diaphragm weakness or paralysis. In combination with cardiac and lung ultrasound, it can detect patients at risk for difficult weaning, predict weaning outcome and help diagnose the cause of weaning failure. A structured ultrasound approach for these patients is presented.

Electronic supplementary material
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Compliance with Ethical Statement

Conflict of interest
None of the authors has any conflict to declare.

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